S18 Table. Results for meta-analysis of African ancestry cohorts from sequence kernel association test (SKAT) association results for previously reported genes for hemoglobin (HGB), hematocrit (HCT), or white blood cell count (WBC) using TOPMed freeze 5b, Haplotype Reference Consortium (HRC), and 1000G phase

3 as imputation reference panels.

| Gene | ENSG ID | Chr:Start-End | Trait | Reference | # Variants | SKAT P-value | N | Previous ancestry | Previous test type | Reference |
|------------|------------------|--------------------------|--------|---------------------|----------------|-------------------|----------------|-------------------|--------------------|-----------|
| HFE | ENSG0000010704 | 6: 26087281- 26098343 | HGB | TOPMed freeze 5b | 27 | 2.63E-05 | 20822 | EU+AA | Burden | (17) |
| | | | | 1000G | 8 | 1.96E-05 | 20822 | | | |
| | | | | HRC | 8 | 7.25E-05 | 20822 | | | |
| MRPL43 | ENSG00000055950 | 10:100969458 - | HGB | TOPMed | 19 | 0.001 | 20822 | EU | Burden | (17) |
| | | 100987515 | | freeze 5b | | | | | | |
| | | | | 1000G | 5 | 0.013 | 15244 | | | |
| | | | | HRC | na | na | na | | | |
| ММАСНС | ENSG00000132763 | 1:45500053 - | HGB | TOPMed | 26 | 0.008 | 20822 | EU | SKAT | (17) |
| | | 45513382 | | freeze 5b | | | | | | |
| | | | | 1000G | 5 | 0.002 | 20822 | | | |
| | | | | HRC | na | na | na | | | |
| EPO | ENSG00000130427 | 7: 00720800- | HGB | TOPMed | 16 | 0.005 | 20822 | EA+HA+AA | SKAT | (21) |
| | | 100723700 | | freeze 5b | | | | | | |
| | | | | 1000G | 11 | 0.012 | 15021 | | | |
| | | | | HRC | 10 | 0.010 | 20822 | | | |
| PKLR | ENSG00000143627 | 1:155289293- | HGB | TOPMed | 27 | 0.038 | 20822 | Multi | SKAT | (6) |
| | | 155301434 | | freeze 5b | | | | | | |
| | | | | 1000G | 5 | 0.108 | 1749 | | | |
| | | | | HRC | 10 | 0.128 | 20822 | | | |
| MMACHC | ENSG00000132763 | 1:45500340- | HCT | TOPMed | 26 | 0.025 | 20818 | EU | SKAT | (17) |
| | | 45509214 | | freeze 5b | | | | | | |
| | | | | 1000G | 5 | 0.008 | 20818 | | | |
| | | | | HRC | na | na | na | | | |
| PKLR | ENSG00000143627 | 1:155289293- | HCT | TOPMed | 27 | 0.025 | 20818 | Multi | SKAT | (6) |
| | | 155301434 | | freeze 5b | - | 0.045 | 15.405 | | | |
| | | | | 1000G | 5 | 0.047 | 17485 | | | |
| | ENIGG0000010504 | < 2<00 72 01 | HOTE | HRC | 10 | 0.093 | 20818 | | D 1 | (15) |
| HFE CXCR2 | ENSG0000010704 | 6:26087281- | HCT | TOPMed | 27 | 0.001 | 20818 | EU+AA | Burden | (17) |
| | | 26098343 | | freeze 5b 1000G | 8 | 0.013 | 20818 | | | |
| | | | | HRC | 8 | 6.88E-04 | | | | |
| | ENSG00000180871 | 2:218125289- | WBC | TOPMed | 21 | 0.88E-04 0.033 | 20818 20827 | EU | Burden | (17) |
| CACK2 | ENSG000001808/1 | 218137253 | WBC | freeze 5b | 21 | 0.033 | 20827 | EU | Burden | (17) |
| | | 210137233 | | 1000G | 4 | 0.120 | 15025 | | | |
| | | | | HRC | 3 | 0.120 | 20826 | | | |
| IQCJ | ENSG00000214216 | 3:158962235- | WBC | TOPMed | 9 | 0.028 | 20827 | EU+HA+AA | SKAT | (21) |
| | E118000000214210 | 159897366 | WBC | freeze 5b |] | 0.026 | 20021 | EUTHATAA | SKAI | (21) |
| | | 137077300 | | 1000G | 2 | 0.077 | 20826 | _ | | |
| | | | | HRC | na | na | na | _ | | |
| S1PR4 | ENSG00000125910 | 19:3172346- | WBC | TOPMed | 33 | 0.045 | 20827 | EU+HA+AA | SKAT | (21) |
| | 21.5000000125710 | 3180332 | ,,,,,, | freeze 5b | 33 | 0.013 | 20021 | DO IIII III | Simil | (21) |
| | | 2100002 | | 1000G | 4 | 0.137 | 13659 | | | |
| | | | | HRC | 7 | 0.066 | 15025 | | | |

| JAK2 | ENSG00000096968 | 9:4984390- | WBC | TOPMed | 37 | 0.026 | 20827 | Multi | VT | (19) |
|------|-----------------|-------------|-----|-----------|----|-------|-------|-------|----|------|
| | | 5128183 | | freeze 5b | | | | | | |
| | | | | 1000G | 13 | 0.008 | 20826 | | | |
| | | | | HRC | 13 | 0.042 | 20826 | | | |
| TAF3 | ENSG00000165632 | 10:7818504- | WBC | TOPMed | 26 | 0.053 | 20827 | EU | VT | (19) |
| | | 8016627 | | freeze 5b | | | | | | |
| | | | | 1000G | 4 | 0.006 | 20826 | | | |
| | | | | HRC | 9 | 0.036 | 20826 | | | |

Previous ancestry column lists the ancestry group in which the significant gene-blood cell trait association signal was previously reported (Multi, multi-ethnic cohort, EU, European, HA, Hispanic/Latino, AA, African American).

Previous test type- statistical test used for previous gene-blood cell trait association (VT, variable threshold, burden, or SKAT).

Association tests were performed using the mmskat function in EPACTS to adjust for kinship, and then p-values were meta-analyzed using metal. Variants were included in an association test if they were annotated as loss of function, high confidence loss of function, missense, protein altering indels, or synonymous variants with fathmm_MKL scores>0.5. Annotation was completed with WGS annotator (WGSA, (22)) using VEP predicted effects for Ensembl(release 83)/GENCODE v24 transcripts and compiled using WGSAParsr version 5.0.4. A variant was included if the minor allele frequency (MAF) was less than 1% in at least one cohort where the estimated r² exceeded 0.8, and the MAF was less than 5% in all cohorts where the estimated r² exceeded 0.8. A variant was only considered for a particular cohort if the estimated r² exceeded 0.8 in that cohort, meaning that the number of variants was allowed to differ by cohort. The number of variants in the table above is the total number of variants included in any SKAT test in the meta-analysis.

All white blood cell results are adjusted for Duffy variant rs2814778. G6PD is not included because we only considered autosomal variants in this study.